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June 2, 2005

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TSCA Document Control Office (7407) EPA East Bldg, Room 6428 1201 Constitution Ave NW Washington DC 20460

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Attention:

TSCA Section 8(e) Coordinator

Re:

Methyl Oxirane (CASRN 75-56-9)

Dear Sir or Madam:

The American Chemistry Council's Propylene Oxide/Propylene Glycol (PO/PG) Panel (Panel), on behalf of its members, <sup>1</sup> is submitting the following information to the EPA pursuant to current guidance issued by EPA indicating EPA's interpretation of Section 8(e) of the Toxic Substances Control Act. The Panel has made no determination as to whether a significant risk of injury to health or the environment is actually presented by the findings.

The test substance methyl oxirane (CASRN 75-56-9) – also known as propylene oxide - was administered by inhalation to groups of 8 male B6C3F1 mice and 8 male F344 rats at 0, 50, 100, 200, and 400 ppm for 20 days at 6 hours/day. Initial microscopic examination of the nasal sections from both species indicate bilateral hyperplasia of transitional epithelium lining the lateral meatus in the proximal nasal airways along with atrophy of the olfactory epithelium lining the dorsal medial meatus at the highest dosing level.

These effects were previously reported in rats following inhalation exposure to propylene oxide (Eldridge et al., 1995; Rios-Blanco et al., 1997; 2003). Mice have not been studied previously. No tumors were induced in the nasal olfactory epithelium in rats or mice following chronic inhalation exposures (NTP, 1985; Lynch et al., 1984; Kuper et al., 1988).

No written report of these initial microscopic evaluations is yet available.

If you have any questions, please contact me at (703) 741-5630 or via email at anne\_lehuray@americanchemistry.com.

Sincerely yours

Anne P. LeHuray, Ph.D.

Manager, Propylene Oxide/Propylene Glycol Panel

Director, CHEMSTAR®

<sup>1</sup> PO/PG Panel members are The Dow Chemical Company, Huntsman Corporation and Lyondell Chemical Company.



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## **References Cited**

- Eldridge, S.R., Bogdanffy, M.S., Jokinen, M.P. and Andrews, L.S. (1995) Effects of propylene oxide on nasal epithelial cell proliferation in F344 rats. *Fundam. Appl. Toxicol.* **27**, 25-32.
- Kuper, C.F., Reuzel, P.G.J., Feron, V.J. and Verschuuren, H.(1988) Chronic inhalation toxicity and carcinogenicity study of propylene oxide in Wistar rats. *Food Chem. Toxicol*, **26**, 159-167.
- Lynch, D.W., Lewis, T.R., Moorman, W.J., Burg, J.R., Groth, D.H., Khan, A., Ackerman, L.J. and Cockrell, B.Y. (1984) Carcinogenic and toxicologic effects of inhaled ethylene oxide and propylene oxide in F344 rats *Toxicol. Appl. Pharmacol.* **76**, 69-84.
- National Toxicology Program (NTP) (1985) Toxicology and carcinogenesis studies of propylene oxide in F344/N rats and B6C3F1 mice (Inhalation Studies), NTP TR 267, U.S. Dept. of Health and Human Services, Public Health Service, National Institutes of Health, Research Triangle Park, N.C., USA.
- Ríos-Blanco, M.N., Plna, K., Faller, T., Kessler, W., Håkansson, K., Kreuzer, P.E., Ranasinghe, A., Filser, J.G., Segerbäck, D. and Swenberg, J.A. (1997) Propylene Oxide: mutagenesis, carcinogenesis and molecular dose. *Mutat. Res.* **380**, 179-197.
- Rios-Blanco, M.N., Ranasinghe, A., Lee, M.S., Faller, T., Filser, J.G., and Swenberg, J.A. (2003). Molecular dosimetry of N7-(2-hydroxypropyl)guanine in tissues of F344 rats after inhalation exposure to propylene oxide. *Carcinogenesis* **24**:1233-1238.